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C. 01 4 5 5	COLUMN CO			
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EPO-Internal, BIOSIS, MEDLINE, WPI Data, EMBASE, PAJ				
C. DOCUM	ENTS CONSIDERED TO BE RELEVANT			
Category °	Citation of document, with indication, where appropriate, of the re	elevant passages	Relevant to claim No.	
X	TIJSTERMAN MARCEL ET AL: "RNA helicase 1-21 MUT-14-dependent gene silencing triggered in C. elegans by short antisense RNAs" SCIENCE (WASHINGTON D C), vol. 295, no. 5555,		1–21	
	25 January 2002 (2002-01-25), pag	ges		
	694-697, XP002257818		•	
	ISSN: 0036-8075			
	abstract page 694 - page 696			
	figure 1			
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	ner documents are listed in the continuation of box C.	Palent family members are listed	n annex.	
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Form PCT/ISA/210 (second sheet) (July 1992)

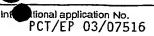


Internation No
PCT/EP 03/07516

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C.(Continua Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
J.,			
X	BOUTLA A ET AL: "Short 5'-phosphorylated double-stranded RNAs induce RNA interference in drosophila" CURRENT BIOLOGY, CURRENT SCIENCE,, GB, vol. 11, no. 22, 13 November 2001 (2001-11-13), pages 1776-1780, XP002965213 ISSN: 0960-9822 table 1	1-21	
X	ELBASHIR S M ET AL: "Analysis of gene function in somatic mammalian cells using small interfering RNAs" METHODS: A COMPANION TO METHODS IN ENZYMOLOGY, ACADEMIC PRESS INC., NEW YORK, NY, US, vol. 26, no. 2, February 2002 (2002-02), pages 199-213, XP002251055 ISSN: 1046-2023 figure 9	1-21	
X	YU JENN-YAH ET AL: "RNA interference by expression of short-interfering RNAs and hairpin RNAs in mammalian cells" PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE UNITED STATES, vol. 99, no. 9, 30 April 2002 (2002-04-30), pages 6047-6052, XP002259803 April 30, 2002 ISSN: 0027-8424 abstract page 6049 - page 6050 page 6047 page 6052 figures 2-4	1-21	
P,X	MARTINEZ JAVIER ET AL: "Single-stranded antisense siRNAs guide target RNA cleavage in RNAi" CELL, vol. 110, no. 5, 6 September 2002 (2002-09-06), pages 563-574, XP002257819 ISSN: 0092-8674 page 569 figure 7	1-21	
Ρ,Χ	SCHWARZ DIANNE S ET AL: "Evidence that siRNAs function as guides, not primers, in the Drosophila and human RNAi pathways" MOLECULAR CELL, vol. 10, no. 3, September 2002 (2002-09), pages 537-548, XP002257820 ISSN: 1097-2765 page 539 - page 541	1-21	



INTERNATIONAL SEARCH REPORT



Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Although claim 21 is directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. X Claims Nos.: 1-21(partially) because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
see PCT/ISA/210 annex
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-21(partially)

Present claims 1-21 relate to the use an extremely large number of possible compounds. Support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT is to be found, however, for only a small proportion of the compounds claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Consequently, the search has been carried out for those parts of the claims which appear to be supported and disclosed, namely those parts relating to RNA interference.

The applicant's attention is drawn to the fact that claims relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-21

Use of single-stranded RNA molecule having a length from 14 - 50 nucleotides wherein at least the 14 - 20 5 most nucleotides are substantially complementary to a target transcript for the manufacture an agent for inhibiting the expression of said target transcript.

2. claims: 22-29

Purified human RISC having a molecular weight of from up to about $150-160~\mathrm{kDa}$.

3. claims: 31-32

An antisense siRNA precursor molecule in the form of a hairpin 25 stem-loop structure comprising 19 to 29 base pairs in stem, wherein at least 14 nucleotides in the stem are substantially complementary to a target transcript.